

ESOPHAGEAL MOTILITY AND PATHOPHYSIOLOGY OF REFLUX DISEASE

OBJECTIVES

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- Mastication and chewing reflex.
- Swallowing process and its stages.
- Nervous initiation of pharyngeal stage of swallowing and its effect on respiration.
- Salivary glands and Contents, Function, Secretion of saliva and control of secretion.
- Types of esophageal peristalsis.
- Esophageal sphincter.
- Function of lower esophageal sphincter.
- Prevention of esophageal reflux by valve like mechanism.

What is the Early Response of the GIT to a Meal? It can be divided into three phases:

Cephalic phase

- Occurs before ingesting food.
- Thinking or anticipating food, smelling or seeing food.
- Aim to prepare the GIT for the meal.
- Stimuli cause an increase in parasympathetic neural outflow to gut leading to enhanced GI secretions (salivary, gastric, pancreatic.. Etc).

Oral phase

- Occurs when ingested food is in the mouth.. (Extra stimulus which is food touching the receptor)
- Same features as cephalic phase but here there is contact between food and GI surface.
- Adding more stimuli from mouth \rightarrow taste.
- The response is similar to the cephalic phase → an increase in parasympathetic neural outflow to gut leading to enhanced GI secretions (salivary,gastric, pancreatic.. Etc).



Lecture Two

- Esophageal phase
- When food is transferred from mouth to esophagus.



3

Where Does Food Digestion Start From? At the mouth (oral cavity)
What is meant by digestion? Breaking down food
What are the types of digestion? Chemical – mechanical
What is the role of the mouth in digestion? Digestion begins in the mouth when food mixes with saliva. Saliva contains an enzyme (amylase) that begins the breakdown of carbohydrates.

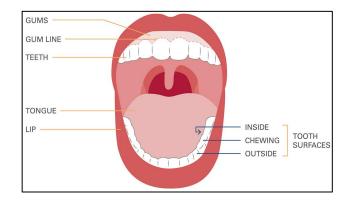


Figure 2-1

Mastication (Chewing):

- **□** Teeth perform cutting and grinding action.
- □ Chewing is largely a *reflex* (5th CN). Functions:
- 1. To lubricate the bolus with salivary secretion.
- 2. To breakdown the bolus to small particles.
- 3. To begin digestion of carbohydrate (by amylase).

| Table 27-1. Functions of Saliva and Chewin | g |
|---|---|
| Disruption of food to produce smaller particles | |
| Formation of a bolus for swallowing | |
| Initiation of starch and lipid digestion | |
| Facilitation of taste | |
| Production of intraluminal stimuli in the stomach | |
| Regulation of food intake and ingestive behavior | |
| Cleansing of the mouth and selective antibacterial action | |
| Neutralization of refluxed gastric contents | |
| Mucosal growth and protection in the rest of the GI tract | |
| Aid in speech | |

Mastication (Chewing)

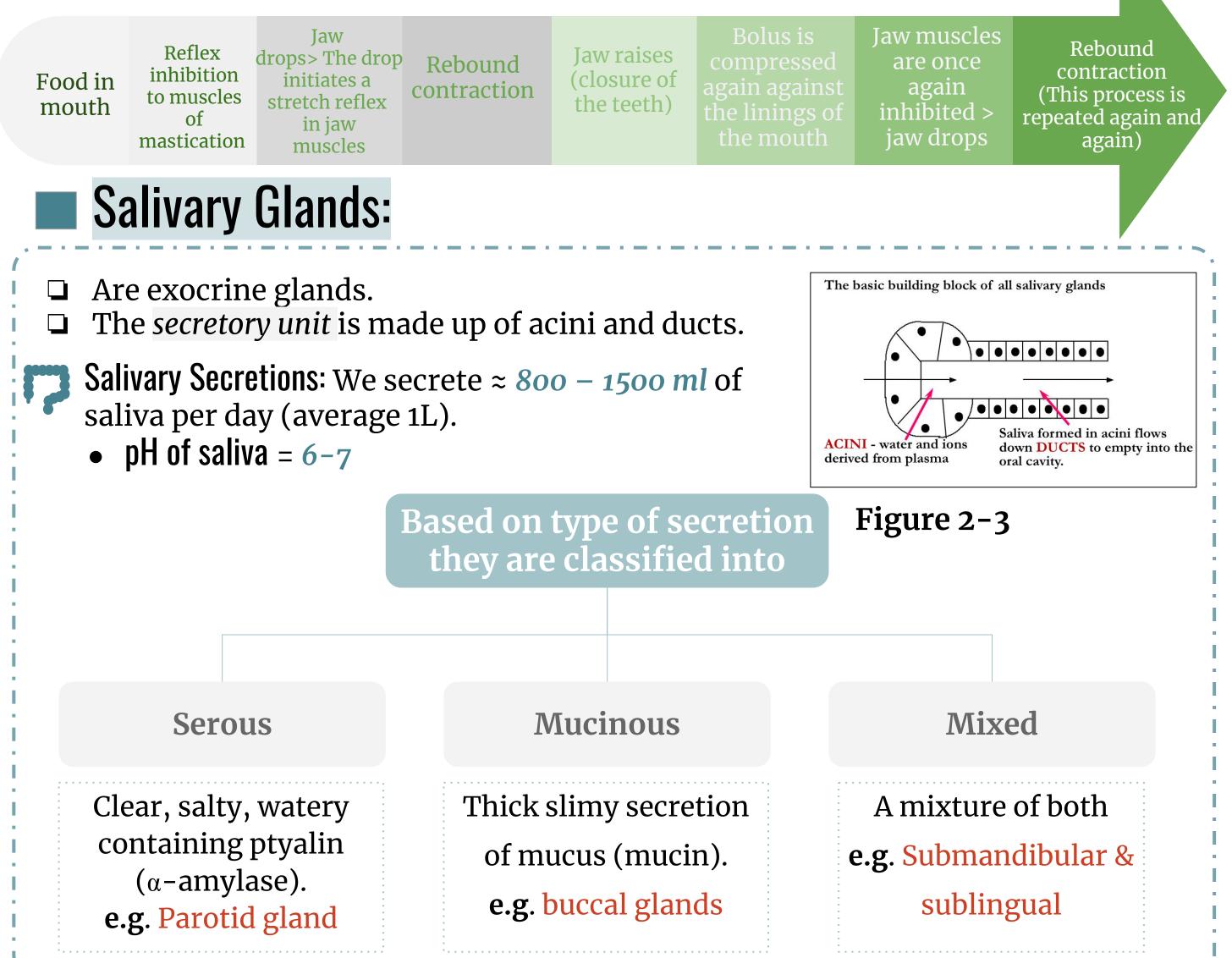
Teeth organization:

Anterior teeth (incisors) for cutting. 2 Posterior teeth (molars) for grinding.

Chewing muscles are innervated by cranial nerve V (trigeminal nerve):

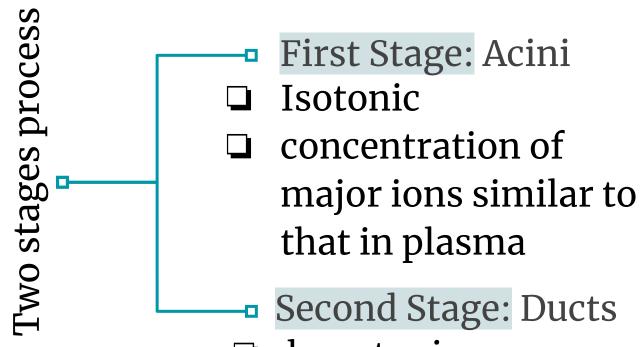
- Masseter
- k Temporalis
- Pterygoid (lateral & medial)
- Taste centers in the brainstem and hypothalamus responsible for rhythmical chewing movements.
- Much of the chewing process is caused by <u>chewing reflex</u> & <u>stretch</u> <u>reflex</u>.

Chewing (Stretch) Reflex





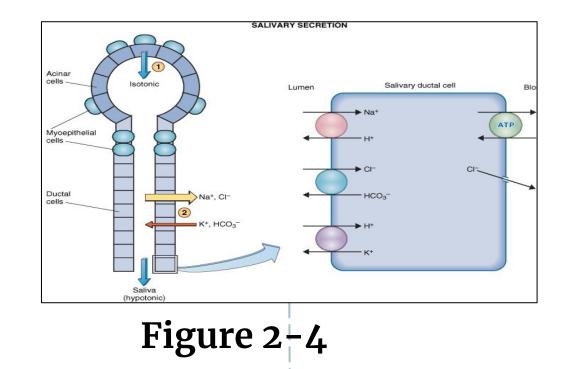
Mechanism of Salivary Secretion



hypotonic

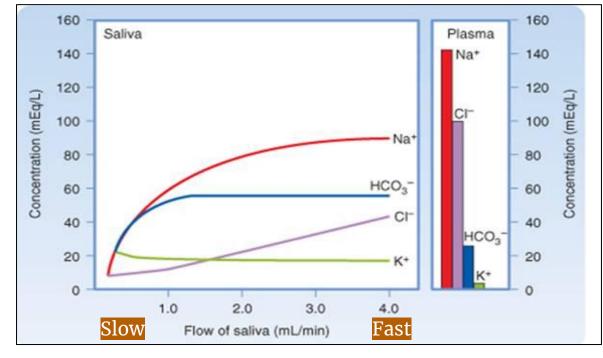
Composition of saliva

- Mucus that serves as a lubricant.
- Amylase: initiates the digestion of starch.
- Lingual lipase: begins digestion of fat.



- Initial saliva is produced by acinar cells & subsequently modified by ductal epithelial cells.
- The fluid secreted from the acini is overall isotonic with the extracellular fluid.
- As the primary secretion (from the acini)
- Electrolytes (Na⁺,Cl⁻, K⁺, HCO₃⁻): moisten food.
- Sodium and chloride ions are usually at a lower concentration than the plasma (hypotonic).
- Potassium, calcium and iodide ions are usually at a higher concentration than plasma (hypertonic)
- Proteins & Enzymes.

flows through the ducts, two major active transport processes take place that markedly modify the ionic composition of the fluid in the saliva. Na⁺ ions are actively reabsorbed from all the salivary ducts and K⁺ ions are actively secreted in exchange for the sodium. Cl[−] ions are passively reabsorbed. HCO₃⁻ ions are secreted by the ductal epithelium into the lumen.



What happens to the composition of saliva in cases of increased flow of secretion?

When the production of saliva is stimulated, flow exceeds the ductal cells maximum rate of modification and so the acinar secretion is modified less (Less hypotonic than resting saliva).

Figure 2-5

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Regulation of Salivary Secretion:

□ Exclusively neural

Increased secretion of salivary glands in response to a meal is a reflex, two types of salivary reflexes;

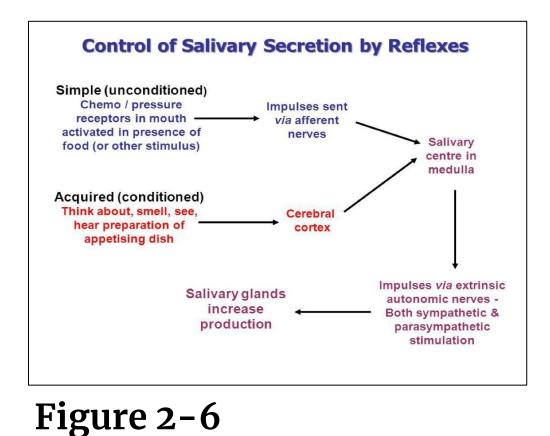
Unconditioned (born with it).



Conditioned reflex (learnt by experience).

Lecture Two

Salivation may occur in response to reflexes originating in the stomach and upper small intestine. E.g. Nauseous, presence of irritating food



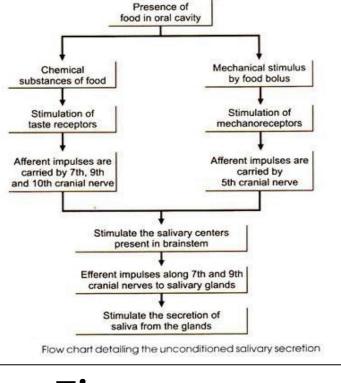
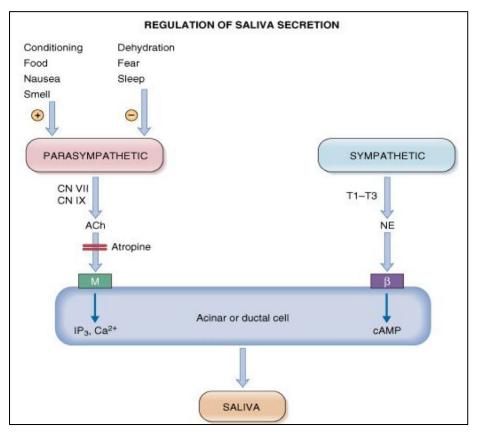


Figure 2-7

Nerve supply of salivary glands

| Sympathetic nerves | Parasympathetic nerves |
|--------------------|------------------------|
| | |

| Originate in the <mark>superior cervical ganglion</mark> and reach the 3 pairs of salivary glands through blood vessels | Originate in the superior & inferior salivary nuclei in brain stem. |
|--|--|
| Functions:- → Act on mucous cells and produce small amount of viscous secretion. → Cause vasoconstriction. | → Fibers from the superior salivary nucleus leave in VII cranial nerve supply both submandibular and sublingual glands. → Fibers from the inferior salivary nucleus leave the medulla in IX cranial nerve supply the parotid gland. |



Stimulation of both sympathetic and parasympathetic nerves stimulates salivary secretion.

Figure 2-8

The Esophageal Phase:

Esophagus:

Collapsible muscular tube that conveys food from pharynx to stomach (10 inches long).

• Food passes through quickly because of peristalsis.

Swallowing (Deglutition)

- Propels food from mouth to stomach.
- Complicated process since pharynx is a shared space between respiration & swallowing.
- Swallowing is initiated voluntarily in the mouth, but thereafter is under involuntary or reflex control. The reflex portion is controlled by the <u>swallowing center</u> in the <u>medulla</u>.
- Food should move without compromising respiration.

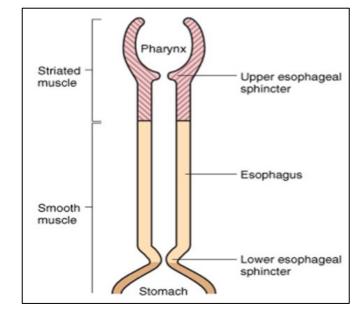


Figure 2-9

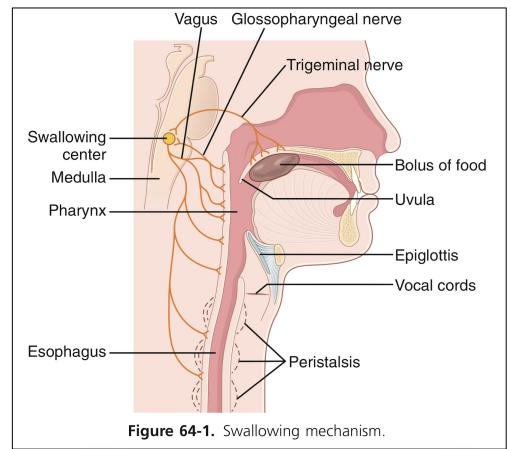


Figure 2-10

Stages of Swallowing:

Moves bolus of food from mouth \rightarrow pharynx Changes: food is squeezed posteriorly into the pharynx by the tongue

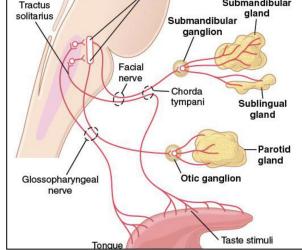


Figure 2-11

2 Pharyngeal Stage of Swallowing (Involuntary)

- ♦ Moves bolus of food from pharynx \rightarrow esophagus.
- Food stimulates touch receptors in pharynx → impulses (Afferent fibers via CN V & IX) → to swallowing center in the brain stem → Motor efferent via CN V, IX, X, & XII → initiate a series of autonomic pharyngeal muscle contractions as follows: (continued on the next page)

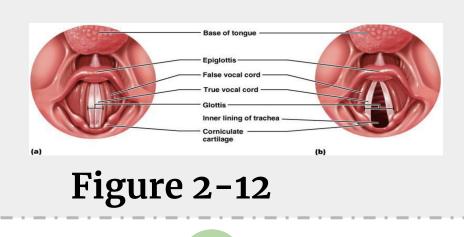
2 Pharyngeal stage of Swallowing (Involuntary)

Soft palate & uvula → pulled upward to close off the nasopharynx Palatopharyngeal folds (on each side of the pharynx) → pulled medially to approximate each other.

→ form a sagittal slit

→ food pass through it to the posterior pharynx.

- **larynx** → pulled upward and anteriorly (by the neck muscles).
- Vocal cords → close
- Epiglottis → bent over the airway as larynx is lifted.
- All these effects prevent food from going into the nose and trachea.
- Destruction of the vocal cords or the muscle that approximate them can cause strangulation.



- **Esophagus** → opens due to the upward movement of the larynx.
- **Upper esophageal sphincter** → relaxes
- **Respiration** → inhibited (the swallowing

The entire muscular wall of the pharynx → contracts (superior, middle, then inferior parts) propelling the food by a wave of **peristalsis**.

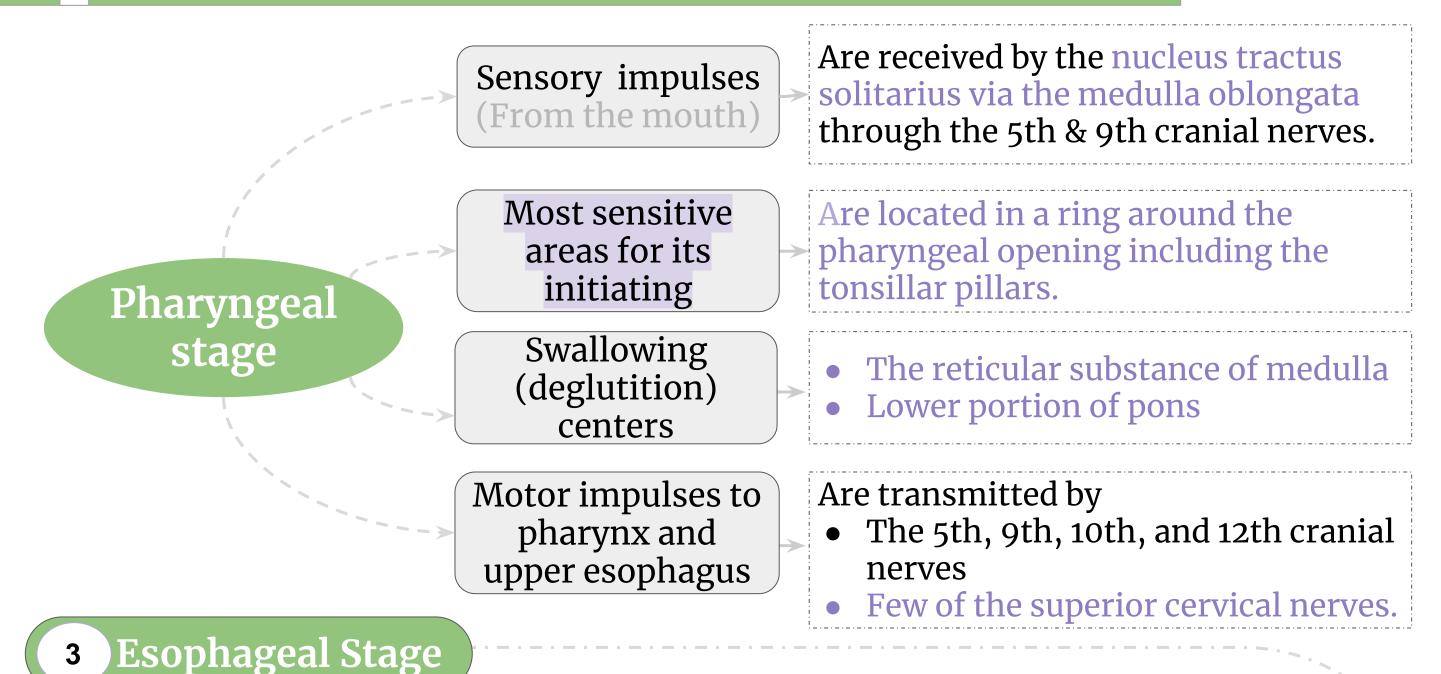
center inhibits the respiratory center in the medulla during the swallowing cycle)

Summary of Pharyngeal Stage of Swallowing:

The pharyngeal stage of swallowing is a reflex act initiated by: The voluntary movement of food into the back of the mouth which stimulates involuntary pharyngeal sensory receptors to elicit the swallowing reflex.

- The trachea is closed.
- The esophagus is opened.
- A fast peristaltic wave initiated by the nervous system of the pharynx forces the bolus of food into the upper esophagus (time of process is < 2 seconds).

ESOPHAGEAL MOTILITY AND PATHOPHYSIOLOGY OF REFLUX DISEASE Lecture Two



- Move food rapidly from pharynx \rightarrow stomach.
- It's controlled partly by the swallowing reflex and partly by the enteric nervous system (ENS). (Myenteric plexus & vagus)
- In case of vagotomy¹ \rightarrow enteric nervous system takes over.
- When bolus of food passes through the upper esophageal sphincter, the swallowing reflex closes the sphincter so food cannot reflux into the pharynx.
- Musculature:
- ★ The pharyngeal wall & upper 1/3 of esophagus: striated muscles, innervated by 10th & 9th cranial nerves.
- ★ Lower two thirds of the esophagus: smooth muscle, controlled by the vagus through connections with the esophageal myenteric nervous system.

Primary peristalsis²

Continuation of the peristaltic wave that started in the pharynx (to push bolus down)

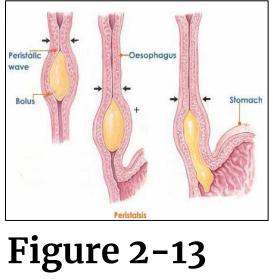
• It passes from the pharynx to the stomach in 8–10 sec

Receptive relaxation³

A wave of relaxation (of LES & stomach) that travels along the myenteric plexus ahead of peristaltic wave. → Allows LES & stomach to prepare to receive the food bolus.

Secondary peristalsis²

- It occurs if the primary peristaltic wave fails to move the food to the stomach
- Starts at the point of esophageal distention by retained food → will continue until all the food is emptied into the stomach.



FOOTNOTES

- A surgical operation in which one or more branches of the vagus nerve are cut, typically to reduce the rate of gastric secretion.
- Nerve pathway of peristalsis: Esophageal distension by bolus → Stimulates mechanoreceptors located on the sensory neurons that lie beneath the epithelium → Sensory neurons send afferent fibers through the vagus nerve to NTS → NTS sends axons to dorsal vagal nucleus (remember this is the autonomic nucleus of the vagus nerve that sends efferents to the GIT) → Axons of dorsal vagus nucleus (vagus efferent fibers) stimulates excitatory motor neurons and inhibitory motor neurons of myenteric plexus → Excitatory motor neurons cause contraction above distension through ACh, inhibitory motor neurons cause relaxation of the segment below the bolus through NO.

Esophageal Stage of Swallowing

The Upper Esophageal Sphincter (UES)

- Formed of skeletal muscle but is not under voluntary control
- located at the lower end of pharynx
- guards the entrance into the esophagus.

• It prevents :

- esophageal air insufflation¹ during negative intrathoracic pressure events(eg: inspiration.)
- esophagopharyngeal/laryngeal reflux during esophageal peristalsis.
- It relaxes during swallowing for about 1 sec → allowing the bolus to be forced through the relaxed UES.

The Lower Esophageal (Gastroesophageal) Sphincter

- Formed by circular muscles
- Extends <u>3cm</u> above its junction with stomach
- Normally remains tonically constricted → Helps to prevent reflux of gastric juice.
- Valvelike mechanism of short portion of esophagus that extend slightly into the stomach also helps in preventing reflux.
- Relaxes ahead of esophageal peristaltic wave "receptive relaxation" → emptying of the propelled food into the stomach.
- Failure of the sphincter to relax will result in <u>achalasia</u>.



- prevents reflux of gastric secretions into the lower portion of the esophagus.
- a short portion of the esophagus extends slightly into the stomach & caves the esophagus inward (closure of the lower esophagus) in response to increased intra-abdominal pressure.

→ Prevents the high pressure in the stomach from forcing its contents into the esophagus.

• Resting pressure (15-30 mmHg).

The diaphragm wraps around the esophagus at the level of lower esophageal sphincter (LES).
 → contraction of the diaphragm helps to increase the pressure at the LES during inspiration.

Vasoactive intestinal peptide (VIP).

• The gastrin hormone, released from the stomach by food, contracts LES.

- Secretin and cholecystokinine (CCK), are released from the upper small intestine, relax the LES.
- Contraction of the circular musculature of the sphincter is regulated by:
- Nerves
- Hormones
- Neurotransmitter.
- Between swallows, tonic vagal cholinergic impulses maintain contraction to keep the sphincter closed.

FOOTNOTES

1. Esophageal insufflation: When inspiration occurs, the lung inflates as air flows into the lung and the intrathoracic pressure becomes negative, this creates a pressure gradient for air to flow into the respiratory tract, but this can affect the esophagus due to the pharynx serving a respiratory function. When air puffs touch the nasopharynx neuronal signals causes the UES to close, preventing the esophagus and stomach from being filled with air.

Achalasia:

A condition due to high resting pressure at the LES that fails to relax during swallowing. As a result, food transmission from the esophagus into the stomach is prevented.

- Causes: pathology of or absence of the myenteric plexus containing VIP & NO in the lower third of esophagus.
- The musculature of the lower esophagus instead remains contracted and the myenteric plexus has lost the ability to transmit a signal to cause relaxation of the LES.
- As achalasia gets worse, the esophagus gradually enlarged as food collects within it.
- Food becomes putridly infected during the long periods of esophageal stasis causing ulceration of the esophageal mucosa, severe substernal pain or even rupture & death.

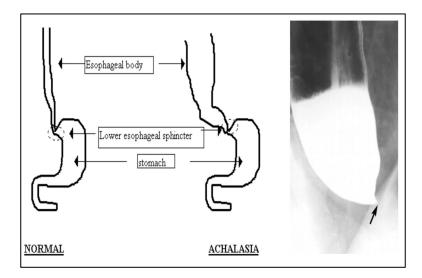


Figure 2-14

Gastroesophageal Reflux Disease (GERD) "Incompetence of the LES"

Incompetence of the lower esophageal sphincter allows reflux of gastric contents into the esophagus.

It may result from a generalized loss of intrinsic sphincter tone or from recurrent inappropriate transient relaxations triggered by gastric distention. Contributing factors are:

- weight gain
- fatty foods
- caffeinated or carbonated beverages
- alcohol
- tobacco smoking
- drugs (as anticholinergics, antihistamines, calcium channel blockers, progesterone and nitrates).

FURTHER READINGS

1. Nerve pathways of peristalsis: Note how the sensory neurons lie beneath the epithelium in Figure 2–14, those sensory neurons contain mechanoreceptors, chemoreceptors, both of which can initiate peristalsis. That's why the presence of acids in the esophagus might initiate peristalsis! But mostly mechanoreceptors sense distension and initiate peristalsis. Peristalsis is aided by gravity but not dependent on it, since you can swallow food upside-down.

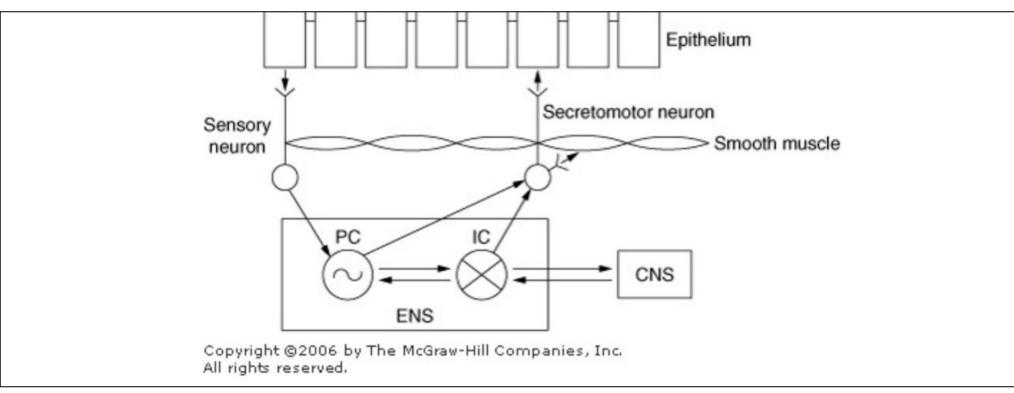


Figure 2-15

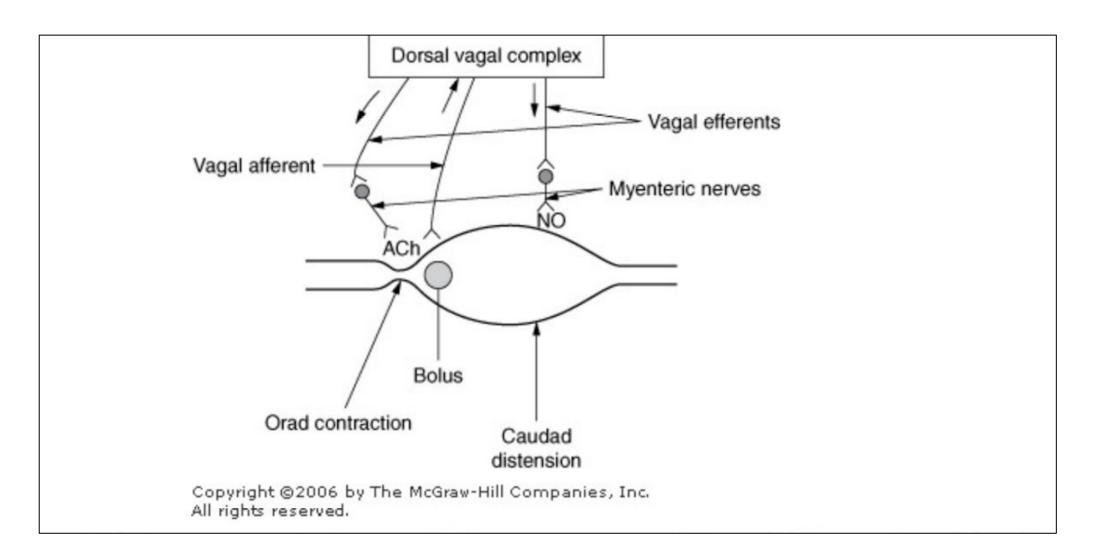


Figure 2-16

Nerve pathway of peristalsis: Esophageal distension by bolus \rightarrow Stimulates mechanoreceptors located on the sensory neurons that lie beneath the epithelium \rightarrow Sensory neurons send afferent fibers through the vagus nerve to NTS \rightarrow NTS sends axons to dorsal vagal nucleus (remember this is the autonomic nucleus of the vagus nerve that sends efferents to the GIT) \rightarrow Axons of dorsal vagus nucleus (vagus efferent fibers) stimulates excitatory motor neurons and inhibitory motor neurons of myenteric plexus \rightarrow Excitatory motor neurons cause contraction above distension by bolus through ACh, inhibitory motor neurons cause relaxation of the segment below the bolus through NO \rightarrow Bolus moves in caudad direction.

QUIZ



- 1. Which of the following physiological process assists in clearing a large food bolus from the esophagus?
- A) Migrating myoelectric complex
- B) Receptive relaxation
- C) Reverse peristalsis
- D) Secondary peristalsis
- 2. The process of mastication is initiated by:
- A) Stretch reflex
- B) Jaw drop
- C) Food bolus inhibiting jaw muscles
- D) Jaw muscles contraction
- 3. Which ONE of the following is most correct about sympathetic stimulation over salivary glands?
- A) Decreases salivary secretion initially
- B) Increase watery secretion of saliva
- C) Decreases salivary secretion when vasoconstriction superimposes
- D) Increases salivary secretion
- E)
- 4. At low flow rate, saliva in the mouth is?
- A) Isotonic, acidic and low in K⁺
- B) Hypotonic, acidic and low in K⁺
- C) Hypertonic, alkaline and rich in Na⁺
- D) Hypotonic, slightly acidic and rich in K⁺
- 5. Which of the following is a mechanism which keeps the valve (of the Gastroesophageal Sphincter) closed.
- A) Valve-like Closure mechanism
- B) Diaphragm helping mechanism
- C) Hormonal and neural mechanism
- D) all of the above

SHORT ANSWER QUESTIONS

Q1: How does the swallowing reflex function?

Q2: What is achalasia?

Q3: List 3 ions in saliva, and their concentration compared to plasma.

- The swallowing reflex center is located in the medulla oblongata. Generator neurons are situated within a primary sensory relay, that is, the nucleus tractus solitarii. which is transmitted by the 5th, 9th, 10th, and 12th cranial nerves and few of the superior cervical nerves.
- 2. A condition due to high resting pressure at the LES that fails to relax during swallowing. As a result, food transmission from the esophagus into the stomach is prevented.
- 3. Na (Low), K (High) and Cl (Low).

ANSWER KEY: D, A, C, D, D



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REFERENCES

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